



UNITED STATES PATENT AND TRADEMARK OFFICE

UNITED STATES DEPARTMENT OF COMMERCE
United States Patent and Trademark Office
Address: COMMISSIONER FOR PATENTS
P.O. Box 1450
Alexandria, Virginia 22313-1450
www.uspto.gov

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/006,116	12/06/2001	Kevin P. Baker	GNE.2830P1C15	8110
35489	7590 04/05/2005		EXAMINER	
	HRMAN WHITE & MCA	MCKELVEY, TERRY ALAN		
MENLO PARK, CO 94025-3506			ART UNIT	PAPER NUMBER
·			1636	
			DATE MAILED: 04/05/2005	

Please find below and/or attached an Office communication concerning this application or proceeding.

	Application No.	Applicant(s)				
	10/006,116	BAKER ET AL.				
Office Action Summary	Examiner	Art Unit				
	Terry A. McKelvey	1636				
The MAILING DATE of this communication	n appears on the cover sheet with	the correspondence address				
Period for Reply						
A SHORTENED STATUTORY PERIOD FOR R THE MAILING DATE OF THIS COMMUNICATI - Extensions of time may be available under the provisions of 37 C after SIX (6) MONTHS from the mailing date of this communicatio - If the period for reply specified above is less than thirty (30) days, - If NO period for reply is specified above, the maximum statutory p - Failure to reply within the set or extended period for reply will, by Any reply received by the Office later than three months after the earned patent term adjustment. See 37 CFR 1.704(b).	ON. FR 1.136(a). In no event, however, may a repon. a reply within the statutory minimum of thirty earlod will apply and will expire SIX (6) MONTI statute, cause the application to become ABA	oly be timely filed (30) days will be considered timely. HS from the mailing date of this communication. NDONED (35 U.S.C. & 133).				
Status						
1) Responsive to communication(s) filed on	12/21/04.					
· _ ·						
	3) Since this application is in condition for allowance except for formal matters, prosecution as to the merits is					
	closed in accordance with the practice under Ex parte Quayle, 1935 C.D. 11, 453 O.G. 213.					
Disposition of Claims						
4)⊠ Claim(s) <u>28-32</u> is/are pending in the appli	cation					
4a) Of the above claim(s) is/are withdrawn from consideration.						
5) Claim(s) is/are allowed.						
6)⊠ Claim(s) <u>28-32</u> is/are rejected.	<u> </u>					
7) Claim(s) is/are objected to.						
8) Claim(s) are subject to restriction a	and/or election requirement.					
Application Papers	·					
9) The specification is objected to by the Exa	miner					
10) ☐ The drawing(s) filed on is/are: a) ☐ accepted or b) ☐ objected to by the Examiner.						
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).						
Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).						
11) The oath or declaration is objected to by the						
Priority under 35 U.S.C. § 119						
<u> </u>	reign priority under 35 LLS C &	119(a) (d) or (f)				
12) Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f). a) All b) Some * c) None of:						
1. Certified copies of the priority documents have been received.						
2. Certified copies of the priority docur		nlication No				
3.☐ Copies of the certified copies of the	* '					
application from the International Be		ooowod w waa waaana olago				
* See the attached detailed Office action for	• • • • • • • • • • • • • • • • • • • •	eceived.				
Attachment(s)	,, []					
 Notice of References Cited (PTO-892) Notice of Draftsperson's Patent Drawing Review (PTO-94) 		mmary (PTO-413) Mail Date				
3) Information Disclosure Statement(s) (PTO-1449 or PTO/S Paper No(s)/Mail Date 12/21/04.		ormal Patent Application (PTO-152)				

Art Unit: 1636

DETAILED ACTION

All objections and rejections not repeated in the instant Action have been withdrawn due to applicant's response to the previous Action.

The text of those sections of Title 35, U.S. Code not included in this action can be found in a prior Office action.

Upon another review of the asserted utilities set forth in the specification, it was determined that the following new rejections are appropriate. Also, a new art rejection is set forth.

Claim Rejections - 35 USC § 101 and 35 USC § 112, First

Paragraph

35 U.S.C. 101 reads as follows:

Whoever invents or discovers any new and useful process, machine, manufacture, or composition of matter, or any new and useful improvement thereof, may obtain a patent therefor, subject to the conditions and requirements of this title.

The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

Art Unit: 1636

Claims 28-32 are rejected under 35 U.S.C. 101 because the claimed invention is not supported by either a specific and substantial asserted utility or a well established utility.

The claims are directed to antibodies that bind isolated polypeptides comprising SEQ ID NO:194, referred to as PRO1303 in the specification, and polypeptides that have 80% or higher amino acid sequence similarity. The specification does not disclose that PRO1303 has significant homology to other, prior art proteins. The instant specification does not disclose any additional information regarding PRO1303 such as subcellular location, timing of regulation during cellular differentiation, which hormones or transcription factors regulate PRO1303, and what physiological significance is possessed by PRO1303. utility of anti-PRO1303 antibodies is linked to the utility of the PRO1303 polypeptide itself because the specification asserts that anti-PRO antibodies can be used in diagnostic assays for PRO, e.g., detecting its expression in specific cells, etc, and used for affinity purification of PRO. Therefore, the antibody has a utility if the protein it binds to, PRO1303, has utility.

The specification also generally asserts that all of the disclosed PRO polypeptides will be useful for a number of purposes; however, none of these asserted uses meet the three-

Application/Control Number: 10/006,116 Page 4

Art Unit: 1636

pronged requirement of 35 USC 101 regarding utility, namely, that the asserted utility be credible, specific, and substantial. The asserted utilities will each be addressed in turn.

- 1. The PRO polypeptide can be used to isolate other polypeptides to which it binds. This asserted utility is not specific or substantial. Since the same can be done with any polypeptide, the asserted utility is not specific to the claimed PRO1303 polypeptide. Furthermore, since the specification does not disclose how PRO1303 or its binding partners can be used, significant further research would be required of the skilled artisan to identify and reasonably confirm a real world context of use because none are set forth simply by binding another protein.
- 2. The PRO polypeptide can be used as a molecular marker. This asserted utility is not specific since the same can be done with any polypeptide and thus is not specific to the PRO1303 polypeptide.
- 3. The PRO polypeptide can be used in tissue typing. The asserted utility is not specific or substantial. With the exception of a few housekeeping genes, all polypeptides have a tissue specific pattern of expression, and thus virtually any

polypeptide can be used in tissue typing. Thus, the asserted utility is not specific to PRO1303.

- 4. The PRO polypeptide can be used in therapy. This asserted utility is not specific or substantial. Since a defect in any polypeptide is likely to cause a disease of some sort, every polypeptide is a target for drug development. Thus, the asserted utility is not specific to the claimed PRO1303 polypeptide. Furthermore, the specification does not disclose a nexus between any specific disease state and a change in amount or form of PRO1303. Significant further research would have to be conducted to identify such a nexus and to thus identify and confirm a real world context of use. Therefore, the asserted utility is not substantial.
- 5. The PRO polypeptide can be used to identify agonists or antagonists. Since the same can be done with any polypeptide, the asserted utility is not specific to the claimed PRO1303 polypeptide. Furthermore, since no activity has been assigned to PRO1303, the assay cannot be conducted until the specific biological activities of PRO1303 are determined empirically. Therefore, the asserted utility is also not substantial.

The specification also discloses that DNA encoding for PRO1303 tested positive in a gene amplification assay, with the DNA being amplified over 2-fold in primary lung tumors and colon

Art Unit: 1636

The utilities asserted based upon this positive result are use as diagnostic markers for determining the presence of tumor cells in lung and/or colon tissue samples and utility in cancer therapy and screening for cancer therapeutics. Even though the DNA encoding PRO1303 has diagnostic utility based upon these results, the PRO1303 polypeptide (and thus the corresponding anti-PRO1303 antibody) does not for the following reasons. The increased copy number of DNA does not provide a readily apparent use for the polypeptide because there is no information regarding level of expression, activity, or role in cancer. Increased copy number of DNA in a cancer or transformed cell does not necessarily result in increased level of expression of the polypeptide, as shown by Konoka et al and Pennica et al. Konopka et al teach that: "Protein expression is not related to amplification of the abl gene but to variation in the level of bcr-abl mRNA produced from a single Ph1 template." (abstract). Pennica et al teach that: "In contrast, WISP-2 mapped to human chromosome 20q12-20q13 and its DNA was amplified, but RNA expression was reduced (2- to >30 fold) in 79% of the tumors." (abstract). These references thus show that even if amplification of a gene occurs in a tumor cell, it does not mean that the mRNA or protein expressed from the gene is also amplified and thus usable as a diagnostic marker for

Art Unit: 1636

cancer. Since the protein is not necessarily overexpressed in cancer cells, then there is no substantial utility in using the protein for cancer therapy or screening for cancer therapeutics.

A substantial utility, by definition is a utility that defines "real world" use, and a utility that requires or constitutes carrying out further research to identify or reasonably confirm a "real world" context of use is not a substantial utility. In the instant case, the amplification of the DNA encoding PRO1303 is, at most, an interesting invitation for further research and confirmation as to whether PRO1303 protein itself is overexpressed or whether high PRO1303 protein expression plays an important role in cancer (and thus might be usable as a therapeutic target). This further research and experimentation, however, is part of the act of invention, and until it has been undertaken, the claimed invention is not considered to have utility based upon gene amplification in tumors.

The specification also discloses that PRO1303 tested positive as stimulators of glucose and/or FFA (free fatty acid) uptake. The asserted utility based upon this assay result is that the polypeptide would be expected to be useful for the therapeutic treatment of disorders where either stimulation or inhibition of glucose uptake by adipocytes would be beneficial

Art Unit: 1636

for example, obesity, diabetes, or hyper- or hypo-insulinemia. The specification does not specifically assert how antibodies against PRO1303 would be used in any of the suggested treatments. First, the specification does not indicate which asserted utilities correspond specifically to glucose uptake stimulation as opposed to glucose uptake inhibition. Second, the specification does not indicate what, if any of the utilities set forth correspond to stimulation of FFA uptake. Third, the actual assay result is stimulation of glucose and/or FFA uptake, three very different activities (stimulation of glucose uptake only, stimulation of FFA uptake only, and stimulation of uptake of both). Would PRO1303 polypeptides be useful to treat hyper-insulinemia or would it be useful to treat hypo-insulinemia, two opposite conditions? Fourth, it is unclear how increasing uptake of FFA into adipocytes would treat obesity (or thus diabetes). Fabris et al teaches that in obesity, excessive energy storage as fat is mainly due to an imbalance between energy intake and expenditure, and the preferential channeling of excess calories as fat rather than protein or glycogen may play an important role in the development and maintenance of the disease. FFA-induced insulin resistance saves scarce glucose for central nervous system requirements, but this becomes counterproductive in obesity

because it inhibits glucose utilization when there is no need to save it. Glucose and FFA might thus be channeled toward tissues (such as adipose tissue in which insulin sensitivity is maintained or even improved) (page 601, second column). Thus, increase of uptake of FFA and/or glucose into adipocytes does not appear to be a utility for treatment of obesity or diabetes.

Furthermore, the observed differences do not appear to be statistically significant and the cutoff points appear to be arbitrary and there is not obvious scientific basis for them. For example, Santomauro et al. (1999. Diabetes 48:1836-1841) teach that 56.5% decreases in FFA levels are statistically significant and correlated with physiological improvements, but it is not clear from either the prior art or the specification whether 50% decreases are useful (see Table 2 from Santomauro et al.). Note that 50% decreases in plasma insulin do appear to be significant, but it is not clear whether this is due to a doubling of insulin uptake by adipocytes or by other tissues, or whether it is due to changes in the amount of insulin production. Similarly, the observation that 56.5% decreases in circulating FFAs is significant and correlated with physiological improvements does not indicate that a doubling of uptake of FFAs by adipocytes will lead to the same decreases in FFAs. For example, doubling the amount of FFA uptake from 1% to

2% of total circulating FFAs would not be expected to lead to a 56% decrease in circulating FFA levels.

35 USC § 101 specifically requires that the invention must be useful in currently available form, which precludes any further experimentation to establish the utility of the claimed invention. Because the instant specification, as filed, fails to disclose a specific role of PRO1303 in glucose and/or FFA uptake in adipocytes, one would have reasons to conclude that the instant invention was not completed as filed, and, therefore, clearly lacks utility in currently available form.

A substantial utility, by definition, is a utility that defines "real world" use, and a utility that requires or constitutes carrying out further research to identify or reasonably confirm a "real world" context of use is not a substantial utility. In the instant case, the mere fact that the protein encoded by the claimed nucleic acids was "positive" in two assays is at the most, an interesting invitation for further research, experimentation and confirmation as to whether the PRO1303 protein or its corresponding antibody is useful as a treatment for diabetes, obesity, hyper-insulinemia, or hypoinsulinemia. The further research and experimentation, however, is part of the act of invention, and until it has been

Art Unit: 1636

undertaken, the claimed invention is not considered specific or substantial.

Claims 28-32 are also rejected under 35 U.S.C. 112, first paragraph. Specifically, since the claimed invention is not supported by either a credible, specific and substantial asserted utility or a well established utility for the reasons set forth above, one skilled in the art clearly would not know how to use the claimed invention.

Claim Rejections - 35 USC § 102

The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless -

(e) the invention was described in (1) an application for patent, published under section 122(b), by another filed in the United States before the invention by the applicant for patent or (2) a patent granted on an application for patent by another filed in the United States before the invention by the applicant for patent, except that an international application filed under the treaty defined in section 351(a) shall have the effects for purposes of this subsection of an application filed in the United States only if the international application designated the United States and was published under Article 21(2) of such treaty in the English language.

Claims 28-32 are rejected under 35 U.S.C. 102(e) as being anticipated by Ni et al (U.S. Patent No. 6,566,498 B1).

Ni et al teach an isolated human secreted polypeptide consisting of SEQ ID NO:6, which has two regions of 100%

Art Unit: 1636

identity with a polypeptide consisting of SEQ ID NO:194, one 62 amino acids long (the first 62 amino acids) and one about 93 amino acids long (at the C-terminus). These large stretches of perfect identity between the two proteins only at each terminus of the proteins would appear to indicate that the two proteins are likely splice-variants of each other and thus are very likely to have many exterior-exposed epitope domains in common. See the attached sequence comparison. Also, because antigenic epitopes can be as low as 7 amino acids and preferably between 15 and 30 amino acids (Column 20 of Ni et al), many of the antibodies taught by the reference which are directed against the protein of SEQ ID NO:6 would strongly cross-react with and specifically bind to the polypeptide of SEQ ID NO:194. "Specifically binds" is interpreted in the claims as broad as is reasonable in the art, which encompasses antibody binding to a protein with a high affinity, of a level comparable with proteins having the identical epitope. The many stretches of 15 to 30 amino acids in common shows that many epitopes that would generate antibodies that specifically bind are in common between the two proteins. Monoclonal and polyclonal antibodies are taught, as are antibody fragments, labeled antibodies, and humanized antibodies (columns 20-21 and 26).

Art Unit: 1636

Conclusion

No claims are allowed.

Certain papers related to this application may be submitted to Art Unit 1636 by facsimile transmission. The faxing of such papers must conform with the notices published in the Official Gazette, 1156 OG 61 (November 16, 1993) and 1157 OG 94 (December 28, 1993) (see 37 C.F.R. § 1.6(d)). The official fax telephone number for the Group is 571-273-8300. NOTE: If Applicant does submit a paper by fax, the original signed copy should be retained by applicant or applicant's representative. NO DUPLICATE COPIES SHOULD BE SUBMITTED so as to avoid the processing of duplicate papers in the Office.

Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to (571) 272-0547.

Patent applicants with problems or questions regarding electronic images that can be viewed in the Patent Application Information Retrieval system (PAIR) can now contact the USPTO's Patent Electronic Business Center (Patent EBC) for assistance. Representatives are available to answer your questions daily from 6 am to midnight (EST). The toll free number is (866) 217-9197. When calling please have your application serial or patent number, the type of document you are having an image problem

with, the number of pages and the specific nature of the problem. The Patent Electronic Business Center will notify applicants of the resolution of the problem within 5-7 business days. Applicants can also check PAIR to confirm that the problem has been corrected. The USPTO's Patent Electronic Business Center is a complete service center supporting all patent business on the Internet. The USPTO's PAIR system provides Internet-based access to patent application status and history information. It also enables applicants to view the scanned images of their own application file folder(s) as well as general patent information available to the public.

For all other customer support, please call the USPTO Call Center (UCC) at 800-786-9199.

Any inquiry concerning rejections or objections in this communication or earlier communications from the examiner should be directed to Terry A. McKelvey whose telephone number is (571) 272-0775. The examiner can normally be reached on Monday through Friday, except for Wednesdays, from about 7:30 AM to about 6:00 PM. A phone message left at this number will be responded to as soon as possible (i.e., shortly after the examiner returns to his office).

Art Unit: 1636

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Dr. Remy Yucel can be reached at (571) 272-0781.

Terry A. McKelvey, Ph.D.

Primary Examiner Art Unit 1636

March 20, 2005

En amile

```
e
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          S &
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          KESULT 3
US-109-244-111-6
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          5 1 5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 å
S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 B
                                                                                            ંઠ
                            ક
                                                             ₽
                                                                                                                                                                                                                                  ; TYPE: PRT
; ORGANISM: Homo sapiens
US-09-244-111-6
                                                                                                                                                                                                                                                                                                                                                                                                                                                     GENERAL INFORMATION:
APPLICANT: N1, et al.
TITLE OF INVENTION: Human Serine Protease and Serpin Polypeptides
FILE REFERENCE: PF391
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 6, Application US/09244111 Patent No. 6566498
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            SOPTWARE: FASTSEQ for Windows Version 4.0 SEQ ID NO 6948
LENGTH: 254
                                                                                                                                                      Matches 159;
                                                                                                                                                                             Query Match
Best Local
                                                                                                                                                                                                                                                                                                                 SEQ ID NO 6
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Query Match 94.7%; Score 1301; DB 4; Length 254; Best Local Similarity 100.0%; Pred. No. 1.5e-118; Matches 235; Conservative 0; Mismatches 0; Indels
                                                                                                                                                                                                                                                                                                                               CURRENT APPLICATION NUMBER: US/09/244,111
CURRENT FILING DATE: 1999-02-04
EARLIER APPLICATION NUMBER: 60/073,961
EARLIER FILING DATE: 1998-02-06
NUMBER OF SEQ ID NOS: 13
SOFTWARE: Patentin Ver. 2.0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ORGANISM: Human
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            PRIOR APPLICATION NUMBER: 60/231,498 PRIOR FILING DATE: 2000-09-08
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    PRIOR FILING DATE: 2000-10-20
PRIOR APPLICATION NUMBER: 60/237,768
PRIOR FILING DATE: 2000-10-03
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             PRIOR APPLICATION NUMBER: 60/241,755
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      CURRENT APPLICATION NUMBER: US/09/949,016
CURRENT FILING DATE: 2000-04-14
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              NUMBER OF SEQ ID NOS:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   APPLICANT: VENTER, J. Craig et al.
TITLE OF INVENTION: POLYMORPHISMS IN KNOWN GENES ASSOCIATED
TITLE OF INVENTION: WITH HUMAN DISEASE, METHODS OF DETECTION AND USES THEREOF
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           FILE REFERENCE: CL001307
                                                                                                                                                                                                                                                                                            LENGTH:
                                                                                                                                                                           Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          121
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 61
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         61
13
                                 61 AHCSGSRYWVRLGEHSLSQLDWTEQIRHSGFSVTHPGYLGASTSHEHDLRLLRLRLPVRV 120
                                                                                                                                                                                                                                                                                            162
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              -
                                                                                              1 MGLSIFLLLCVLGLSQAATPKIFNGTECGRNSQPWQVGLFEGTSLRCGGVLIDHRWVLTA 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1 MGLSIFLLLCVLGLSQAATPKIFNGTECGRNSQPWQVGLFEGTSLRCGGVLIDHRWVLTA 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             TSNMVCAGGVPGQDACQGDSGGPLVCGGVLQGLVSWGSVGPCGQDGIPGVYTYIC 235
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            TSSVQPLPLPNDCATAGTECHVSGWGITNHPRNPFPDLLQCLNLSIVSHATCHGVYPGGRI 180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AHCSGSRYWVRLGEHSLSQLDWTEQIRHSGFSVTHPGYLGASTSHEHDLRLLRLRLPVRV 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TSSVQPLPLPNDCATAGTECHVSGWGITNHPRNPFPDLLQCLNLSIVSHATCHGVYPGRI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AHCSGSRYWVRLGEHSLSQLDWTEQIRHSGFSVTHPGYLGASTSHEHDLRLLRLPVRV 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            MGLSIFLLLCVLGLSQAATPKI FNGTECGRNSQPWQVGLFEGTSLRCGGVLIDHRWVLTA 60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TSNMVCAGGVPGQDACQGDSGGPLVCGGVLQGLVSWGSVGPCGQDGIPGVYTYIC 235
                                                                                                                                                      59.0%; ilarity 64.1%; Conservative
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    2000-04-14
                                                                                                                                                    Score 811; DB 4; I Pred. No. 3.8e-71; 0; Mismatches 3;
                                                                                                                                                                                         DB 4; Length 162;
                                                                                                                                                      Indels
                                                                                                                                                      86;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            0
                                                                                                                                                      Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Gaps
                                                                        60
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       180
```

US-09-070-526-2

Sequence 2, Application US/09070526 Patent No. 6100059

GENERAL INFORMATION:
APPLICANT: SOUTHAN, CHRISTY
APPLICANT: CLINKENBEARD, HE
APPLICANT: BURGESS, NICOLA
TITLE OF INVENTION: NO. 610

6100059el HELEN

SOUTHAN, CHRISTOPHER

CORRESPONDENCE ADDRESS: NUMBER OF SEQUENCES: ADDRESSEE:

RATNER & PRESTIA O. BOX 980

```
APPLICANT: O'Brien, Timothy J.
APPLICANT: Underwood, Lowell J.
TITLE OF INVENTION: No. 6642013el Extracell
FILE REFERENCE: D6020CIP2
CURRENT APPLICATION NUMBER: US/09/618,259
CURRENT FILING DATE: 2000-07-18
PRIOR APPLICATION NUMBER: US 09/127,444
PRIOR APPLICATION NUMBER: US 09/127,444
PRIOR FILING DATE: 198-08-21
NUMBER OF SEQ ID NOS: 72
SEQ ID NO 7
 밁
                                 S
                                                                     밁
                                                                                                       5
                                                                                                                                             밁
                                                                                                                                                                             S
                                                                                                                                                                                                                    문
                                                                                                                                                                                                                                                                                          밁
                                                                                                                                                                                                                                                                                                                            S
                                                                                                                                                                                                                                                    S
                                                                                                                                                                                                                                                                                                                                                                                                                                       ; OTHER INFORMATION: Amino acid sequence of TADG-14 protein US-09-618-259-7
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    US-09-618-259-7
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                문
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              5
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                GENERAL INFORMATION:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 7, Application US/09618259 Patent No. 6642013
                                                                                                                                                                                                                                                                                                                                                                   Matches
                                                                                                                                                                                                                                                                                                                                                                                  Query Match
Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TYPE: PRT
ORGANISM: Homo sapiens
FEATURE:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            LENGTH: 260
                                                                                            181 TSNMYCAGGVPGQDACQGDSGGPLVCGGVLQGLVSWGSVGPCGQDGIPGVYTYICKYVDW
 252
                                   241 IRMIM 245
                                                                         193
                                                                                                                                               133
                                                                                                                                                                                121
                                                                                                                                                                                                                                                                                                                                                                 123;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     181
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                241
                                                                                                                                                                                                                    73
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         121 TSSVQPLPLPNDCATAGTECHVSGWGITNHPRNPFPDLLQCLNLSIVSHATCHGVYPGRI 180
                                                                                                                                                                                                                                         62 HCSGSRYWVRLGEHSLSQLDWTEQIRHSGFSVTHPGYLGAST-SHEHDLRLLRLRLPVRV 120
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    95
                                                                                                                                                                                                                                                                                        13 MFLLLLGGAWAGHSRAQEDKVLGGHECQPHSQPWQAALFQGQQLLCGGVLVGGNWVLTAA
                                                                                                                                                                                                                                                                                                                  5 IFLLL---CVLGLSQAATPKIFNGTECGRNSQPWQVGLFEGTSLRCGGVLIDHRWVLTAA
                                                                                                                                             GSKVKPISLADHCTQPGQKCTVSGWGTVTSPRENFPDTLNCAEVKIFPQKKCEDAYPGQI
                                                                                                                                                                               TSSYQPLPLPNDCATAGTECHVSGWGITNHPRNPFPDLLQCLNLSIVSHATCHGVYPGRI 180
                                                                                                                                                                                                                    HCKKPKYTVRLGDHSLQNKDGPEQEIPVVQSIPHPCYNSSDVEDHNHDLMLLQLRDQASL 132
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        TSNMVCAGGVPGQDACQGDSGGPLVCGGVLQGLVSWGSVGPCGQDGIPGVYTYICKYVDW 240
IKKII 256
                                                                       TDGMVCAGSSKGADTCQGDSGGPLVCDGALQGITSWGS-DPCGRSDKPGVYTNICRYLDW
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              IRMIMRNN 248
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     TSNMVCAGGVPGQDACQGDSGGPLVCGGVLQGLVSWGSVGPCGQDGIPGVYTYICKYVDW 154
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            IRMIMRNN 162
                                                                                                                                                                                                                                                                                                                                                                Conservative
                                                                                                                                                                                                                                                                                                                                                                45.9%; Score 630.5;
50.2%; Pred. No. 2.50
tive 24; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         6642013el Extracellular Serine Protease
                                                                                                                                                                                                                                                                                                                                                                                                   DB
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           -PDLLQCLNLSIVSHATCHGVYPGRI
                                                                                                                                                                                                                                                                                                                                                                 93;
                                                                                                                                                                                                                                                                                                                                                                                                   4
                                                                                                                                                                                                                                                                                                                                                                                                 Length
                                                                                                                                                                                                                                                                                                                                                                Indels
                                                                                                                                                                                                                                                                                                                                                                                                 260;
                                                                                                                                                                                                                                                                                                                                                                ហ
                                                                                                                                                                                                                                                                                                                                                                Gaps
                                                                                                           240
                                                                                                                                             192
                                                                                                                                                                                                                                                                                            72
                                                                       251
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           94
```